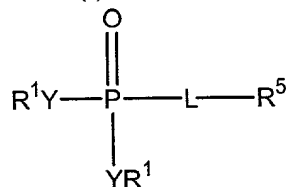
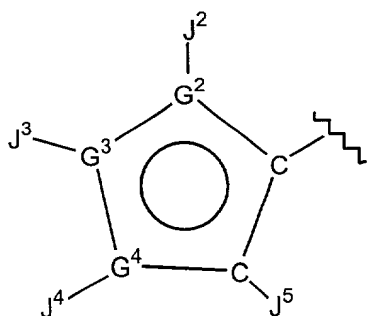


**We claim:**

1. A compound of formula (I):

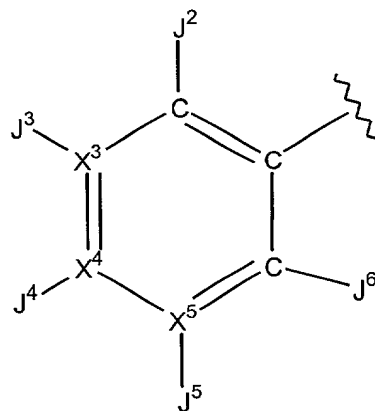


(I)

wherein  $\text{R}^5$  is selected from the group consisting of:

I (a)

and



I (b)

wherein:

 $\text{G}^2$  is selected from the group consisting of C, O, and S; $\text{G}^3$  and  $\text{G}^4$  are independently selected from the group consisting of C, N, O, and S;

wherein a) not more than one of  $\text{G}^2$ ,  $\text{G}^3$ , and  $\text{G}^4$  may be O, or S; b) when  $\text{G}^2$  is O or S, not more than one of  $\text{G}^3$  and  $\text{G}^4$  is N; c) at least one of  $\text{G}^2$ ,  $\text{G}^3$ , and  $\text{G}^4$  is C; and d)  $\text{G}^2$ ,  $\text{G}^3$ , and  $\text{G}^4$  are not all C;

 $\text{X}^3$ ,  $\text{X}^4$ , and  $\text{X}^5$  are independently selected from the group consisting of C and N,wherein no more than two of  $\text{X}^3$ ,  $\text{X}^4$ , and  $\text{X}^5$  may be N;

$\text{J}^2$ ,  $\text{J}^3$ ,  $\text{J}^4$ ,  $\text{J}^5$ , and  $\text{J}^6$  are independently selected from the group consisting of -H,  $-\text{NR}^4_2$ ,  $-\text{CONR}^4_2$ ,  $-\text{CO}_2\text{R}^3$ , halo,  $-\text{S}(\text{O})_2\text{NR}^4_2$ ,  $-\text{S}(\text{O})\text{R}^3$ ,  $-\text{SO}_2\text{R}^3$ , alkyl, alkenyl, alkynyl, alkylenearyl, perhaloalkyl, haloalkyl, aryl, heteroaryl, alkylene-OH,  $-\text{C}(\text{O})\text{R}^{11}$ ,  $-\text{OR}^{11}$ , -

alkylene- $\text{NR}^4_2$ , -alkylene-CN, -CN, -C(S) $\text{NR}^4_2$ , - $\text{OR}^2$ , - $\text{SR}^2$ , - $\text{N}_3$ , - $\text{NO}_2$ , - $\text{NHC(S)NR}^4_2$ , and - $\text{NR}^{18}\text{COR}^2$ ;

L is selected from the group consisting of:

i) a linking group having 2-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -furanyl-, -thienyl-, -pyridyl-, -oxazolyl-, -imidazolyl-, -phenyl-, -pyrimidinyl-, -pyrazinyl-, and -alkynyl-, all of which may be optionally substituted; and

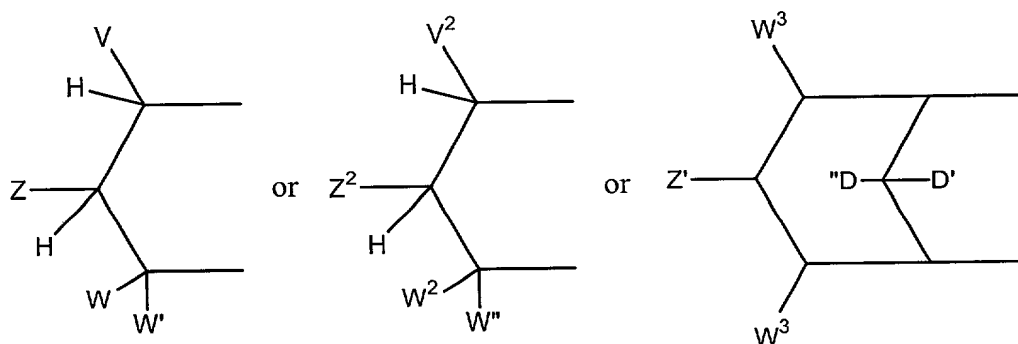
ii) a linking group having 3-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -alkylenecarbonylamino-, -alkyleneaminocarbonyl-, -alkyleneoxycarbonyl-, -alkyleneoxy-, and -alkyleneoxyalkylene-, all of which may be optionally substituted;

Y is independently selected from the group consisting of -O-, and - $\text{NR}^6$ -;

when Y is -O-, then  $\text{R}^1$  attached to -O- is independently selected from the group consisting of -H, alkyl, optionally substituted aryl, optionally substituted alicyclic where the cyclic moiety contains a carbonate or thiocarbonate, optionally substituted arylalkylene-, - $\text{C(R}^2)_2\text{OC(O)NR}^2_2$ , - $\text{NR}^2\text{-C(O)-R}^3$ , - $\text{C(R}^2)_2\text{-OC(O)R}^3$ , - $\text{C(R}^2)_2\text{-O-C(O)OR}^3$ , - $\text{C(R}^2)_2\text{OC(O)SR}^3$ , -alkylene-S-C(O) $\text{R}^3$ , -alkylene-S-S-alkylenehydroxy, and -alkylene-S-S-alkylenehydroxy,

when one Y is - $\text{NR}^6$ -, and  $\text{R}^1$  attached to it is - $(\text{CR}^{12}\text{R}^{13})_n\text{-C(O)-R}^{14}$ , then the other  $\text{YR}^1$  is selected from the group consisting of - $\text{NR}^{15}\text{R}^{16}$ , - $\text{OR}^7$ , and  $\text{NR}^6\text{-(CR}^{12}\text{R}^{13})_n\text{-C(O)-R}^{14}$ ;

or when either Y is independently selected from -O- and - $\text{NR}^6$ -, then together  $\text{R}^1$  and  $\text{R}^1$  are -alkylene-S-S-alkylene- to form a cyclic group, or together  $\text{R}^1$  and  $\text{R}^1$  are



wherein

a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

5 Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC(O)R}^3$ ,  $-\text{CHR}^2\text{OC(S)R}^3$ ,  $-\text{CHR}^2\text{OC(S)OR}^3$ ,  $-\text{CHR}^2\text{OC(O)SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH(aryl)OH}$ ,  $-\text{CH(CH=CR}^2_2)\text{OH}$ ,  $-\text{CH(C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p\text{-OR}^{19}$ , and  $-(\text{CH}_2)_p\text{-SR}^{19}$ ;

or

10 together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

15 W and W' are independently selected from the group of  $-\text{H}$ , alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;

or

20 together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

b)  $\text{V}^2$ ,  $\text{W}^2$  and  $\text{W}''$  are independently selected from the group of  $-\text{H}$ , alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group  
5 containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with  
hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon  
atom that is three atoms from a Y attached to phosphorus;

c)  $Z'$  is selected from the group of  $-\text{OH}$ ,  $-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{OCO}_2\text{R}^3$ , and  
10  $-\text{OC}(\text{O})\text{SR}^3$ ;

$D'$  is  $-\text{H}$ ;

$D''$  is selected from the group of  $-\text{H}$ , alkyl,  $-\text{OR}^2$ ,  $-\text{OH}$ , and  $-\text{OC}(\text{O})\text{R}^3$ ;

each  $W^3$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, aralkyl,  
alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

15  $p$  is an integer 2 or 3;

with the provisos that:

a)  $V$ ,  $Z$ ,  $W$ ,  $W'$  are not all  $-\text{H}$  and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all  $-\text{H}$ ; and

$R^2$  is selected from the group consisting of  $R^3$  and  $-\text{H}$ ;

$R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

20 each  $R^4$  is independently selected from the group consisting of  $-\text{H}$ , alkylene,  
 $-\text{alkylenearyl}$  and aryl, or together  $R^4$  and  $R^4$  are connected via 2-6 atoms, optionally  
including one heteroatom selected from the group consisting of O, N, and S;

$R^6$  is selected from the group consisting of  $-\text{H}$ , lower alkyl, acyloxyalkyl, aryl,  
aralkyl, alkyloxycarbonyloxyalkyl, and lower acyl, or together with  $R^{12}$  is connected via 1-  
25 4 carbon atoms to form a cyclic group;

$R^7$  is lower  $R^3$ ;

each  $R^9$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, aralkyl,  
and alicyclic, or together  $R^9$  and  $R^9$  form a cyclic alkyl group;

$R^{11}$  is selected from the group consisting of alkyl, aryl,  $-\text{NR}^2_2$ , and  $-\text{OR}^2$ ; and

each  $R^{12}$  and  $R^{13}$  is independently selected from the group consisting of H, lower alkyl, lower aryl, lower aralkyl, all optionally substituted, or  $R^{12}$  and  $R^{13}$  together are connected via a chain of 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S, to form a cyclic group;

5 each  $R^{14}$  is independently selected from the group consisting of  $-OR^{17}$ ,  $-N(R^{17})_2$ ,  $-NHR^{17}$ ,  $-SR^{17}$ , and  $-NR^2OR^{20}$ ;

$R^{15}$  is selected from the group consisting of  $-H$ , lower aralkyl, lower aryl, lower aralkyl, or together with  $R^{16}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

10  $R^{16}$  is selected from the group consisting of  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ ,  $-H$ , lower alkyl, lower aryl, lower aralkyl, or together with  $R^{15}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

each  $R^{17}$  is independently selected from the group consisting of lower alkyl, lower aryl, and lower aralkyl, or together  $R^{17}$  and  $R^{17}$  on N is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

15  $R^{18}$  is selected from the group consisting of  $-H$  and lower  $R^3$ ;

$R^{19}$  is selected from the group consisting of  $-H$ , and lower acyl;

$R^{20}$  is selected from the group consisting of  $-H$ , lower  $R^3$ , and  $-C(O)-(lower\ R^3)$ ;

$n$  is an integer from 1 to 3;

20 with the provisos that:

- 1) when  $X^3$ ,  $X^4$ , or  $X^5$  is N, then the respective  $J^3$ ,  $J^4$ , or  $J^5$  is null;
- 2) when L is substituted furanyl, then at least one of  $J^2$ ,  $J^3$ ,  $J^4$ , and  $J^5$  is not  $-H$  or null;
- 3) when L is not substituted furanyl, then at least two of  $J^2$ ,  $J^3$ ,  $J^4$ , and  $J^5$  on formula I(a) or  $J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  on formula I(b) are not  $-H$  or null;
- 25 4) when  $G^2$ ,  $G^3$ , or  $G^4$  is O or S, then the respective  $J^2$ ,  $J^3$ , or  $J^4$  is null;
- 5) when  $G^3$  or  $G^4$  is N, then the respective  $J^3$  or  $J^4$  is not halogen or a group directly bonded to  $G^3$  or  $G^4$  via a heteroatom;
- 6) if both Y groups are  $-NR^6-$ , and  $R^1$  and  $R^1$  are not connected to form a cyclic phosphoramidate, then at least one  $R^1$  is  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ ;
- 30

- 7) when L is -alkylenecarbonylamino- or -alkyleneaminocarbonyl-, then  $X^3$ ,  $X^4$ , and  $X^5$  are not all C;
- 8) when L is -alkeneoxyalkylene-, and  $X^3$ ,  $X^4$ , and  $X^5$  are all C, then neither  $J^3$  nor  $J^5$  can be substituted with an acylated amine;
- 5 9) when  $R^5$  is substituted phenyl, then  $J^3$ ,  $J^4$ , and  $J^5$  is not purinyl, purinylalkylene, deaza-purinyl, or deazapurinylalkylene;
- 10)  $R^1$  can be selected from the lower alkyl only when the other  $YR^1$  is  $-NR^6-C(R^{12}R^{13})_n-C(O)-R^{14}$ ;
- 11) when  $R^5$  is substituted phenyl and L is 1,2-ethynyl, then  $J^3$  or  $J^5$  is not a heterocyclic group;
- 10 12) when L is 1,2-ethynyl, then  $X^3$  or  $X^5$  cannot be N;
- and pharmaceutically acceptable prodrugs and salts thereof.

2. The compounds of claim 1 wherein  $R^5$  is selected from the group consisting of substituted phenyl, substituted pyrrolyl, substituted oxazolyl, substituted thiazolyl, substituted isothiazolyl, substituted pyrazolyl, substituted isoxazolyl, substituted pyridinyl, substituted thienyl, substituted furanyl, substituted pyrimidinyl, and substituted pyridazinyl.

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3. The compounds of claim 1 with the further proviso that when L is -alkyleneoxyalkylene-, and  $R^5$  is substituted thienyl, substituted furanyl, or substituted phenyl, then  $J^3$ ,  $J^4$ , or  $J^5$  is not halo or alkenyl.

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4. The compounds of claim 1 with the further proviso that when L is -alkyleneoxyalkylene-, then  $R^5$  is not substituted thienyl, substituted furanyl, or substituted phenyl.

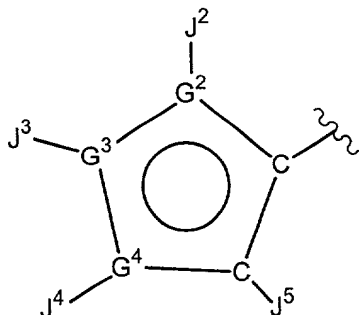
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5. The compounds of claim 1 with the further proviso that when L is -alkyleneoxycarbonyl-, and  $X^3$ ,  $X^4$ , and  $X^5$  are all C, then neither  $J^2$  nor  $J^6$  is a group attached through a nitrogen atom.

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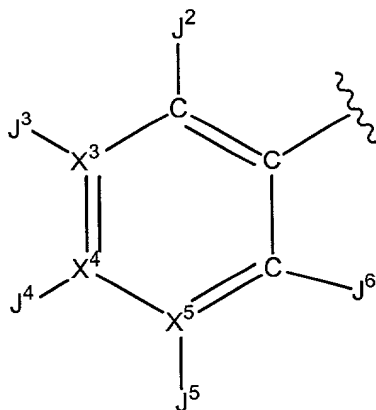
6. The compounds of claim 1 with the further proviso that when L is -alkyleneoxyalkylene- or -alkyleneoxycarbonyl-, then R<sup>5</sup> is not substituted phenyl.

5 7. The compounds of claim 1 wherein R<sup>5</sup> is a compound of formula I(a):



I (a)

8. The compounds of claim 1 wherein R<sup>5</sup> is a compound of formula I(b):



I (b)

10 9. The compounds of claim 1 wherein L is selected from the group consisting of :

- i) 2,5-furanyl, 2,5-thienyl, 2,6-pyridyl, 2,5-oxazolyl, 5,2-oxazolyl, 2,4-oxazolyl, 4,2-oxazolyl, 2,4-imidazolyl, 2,6-pyrimidinyl, 2,6-pyrazinyl, 1,3-phenyl;
- 15 ii) 1,2-ethynyl; and

- iii) a linking group having 3 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -alkylenecarbonylamino-, -alkyleneaminocarbonyl-, -alkyleneoxycarbonyl-, and -alkyleneoxyalkylene-.

10. The compounds of claim 9 wherein L is selected from the group consisting of :

- i) 2,5-furanyl, 2,5-thienyl, 2,6-pyridyl, 2,5-oxazolyl, 5,2-oxazolyl, 2,4-oxazolyl, 4,2-oxazolyl, 2,4-imidazolyl, 2,6-pyrimidinyl, 2,6-pyrazinyl, 1,3-phenyl; and
- ii) 1,2-ethynyl.

11. The compounds of claim 9 wherein L is selected from the group consisting of :

- i) 2,5-furanyl, 2,6-pyridyl, 2,5-oxazolyl, 2,4-imidazolyl, 1,3-phenyl;
- ii) 1,2-ethynyl; and
- iii) a linking group having 3 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -methylenecarbonylamino-, -methyleneaminocarbonyl-, -methyleneoxycarbonyl-, and -methyleneoxymethylene-.

12. The compounds of claim 11 wherein L is selected from the group consisting of 2,5-furanyl, methyleneoxycarbonyl, methyleneoxymethylene, and methyleneaminocarbonyl.

13. The compounds of claim 12 wherein L is 2,5-furanyl.

14. The compounds of claim 1 wherein  $X^4$  and  $X^5$  are C.



15. The compounds of claim 1 wherein  $J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of  $-H$ ,  $-NR^4_2$ ,  $-C(O)NR^4_2$ ,  $-CO_2R^3$ , halo,  $-SO_2NR^4_2$ , lower alkyl, lower alkenyl, lower alkynyl, lower perhaloalkyl, lower haloalkyl, lower aryl, lower alkylaryl, lower alkylene-OH,  $-OR^{11}$ ,  $-CR^2_2NR^4_2$ ,  $-CN$ ,  $-C(S)NR^4_2$ ,  $-OR^2$ ,  $-SR^2$ ,  $-N_3$ ,  $-NO_2$ ,  $-NHC(S)NR^4_2$ ,  $-NR^{18}C(O)R^2$  and  $-CR^2_2CN$ .

16. The compounds of claim 12 wherein  $J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of  $-H$ ,  $-NO_2$ , lower alkyl, lower alkylaryl, lower alkoxy, lower perhaloalkyl, halo,  $-CH_2NHR^4$ ,  $-C(O)NR^4_2$ ,  $-S(O)_2NHR^4$ ,  $-OH$ ,  $-NH_2$ , and  $-NHC(O)R^2$ .

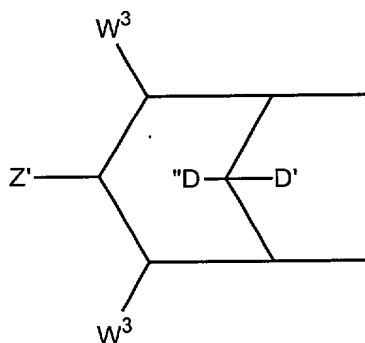
17. The compounds of claim 1, where both Y groups are  $-O-$ .

18. The compounds of claim 1, where both Y groups are  $-NR^6-$ .

19. The compounds of claim 1 where one Y is  $-NR^6-$ , and one Y is  $-O-$ .

20. The compounds of claim 1 wherein each  $YR^1$  is  $-OH$ .

21. The compounds of claim 1 wherein  $R^1$  and  $R^1$  together are



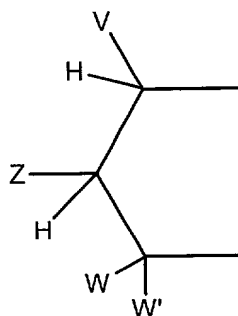
$Z'$  is selected from the group of  $-OH$ ,  $-OC(O)R^3$ ,  $-OCO_2R^3$ , and  $-OC(O)SR^3$ ;

$D'$  is  $-H$ ;

$D''$  is selected from the group of  $-H$ , alkyl,  $-OR^2$ ,  $-OH$ , and  $-OC(O)R^3$ ; and

each  $W^3$  is independently selected from the group consisting of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl.

22. The compounds of claim 1 wherein  $R^1$  and  $R^1$  together are



V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2_2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p-\text{OR}^{19}$ , and  $-(\text{CH}_2)_p-\text{SR}^{19}$ ;  
or

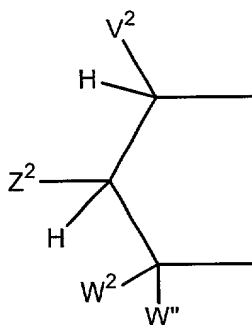
together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

W and  $W'$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;  
or

together W and  $W'$  are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl.

23. The compounds of claim 1 wherein  $R^1$  and  $R^1$  together are



- $V^2$ ,  $W^2$  and  $W'''$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

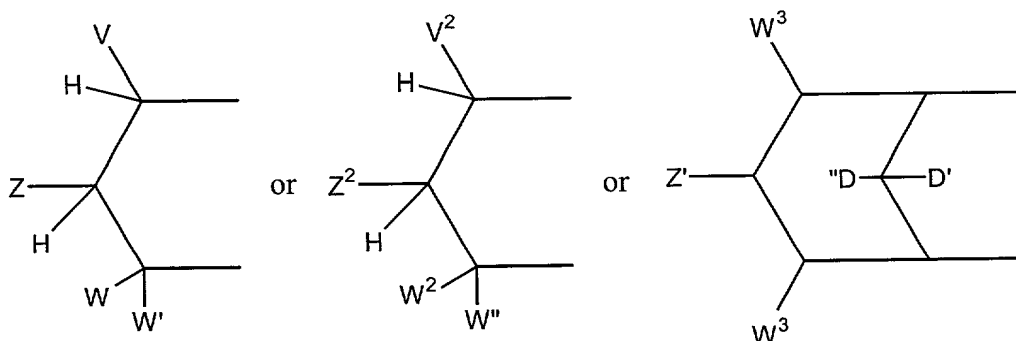
- together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus.

24. The compounds of claim 1 wherein when both Y groups are -O-, then  $R^1$  attached to -O- is optionally substituted aryl.

25. The compounds of claim 1 wherein when both Y groups are -O-, then  $R^1$  is independently selected from the group consisting of optionally substituted aralkyl.

26. The compounds of claim 1 wherein both Y groups are -O-, and at least one  $R^1$  is selected from the group consisting of  $-\text{C}(\text{R}^2)_2-\text{OC}(\text{O})\text{R}^3$ , and  $-\text{C}(\text{R}^2)_2-\text{OC}(\text{O})\text{OR}^3$ .

27. The compounds of claim 1 wherein at least one Y is -O-, and together  $R^1$  and  $R^1$  are



wherein

- 5 a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p\text{-OR}^{19}$ , and  $-(\text{CH}_2)_p\text{-SR}^{19}$ ;  
 10 or

together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

- 15 together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

W and W' are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;

- 20 or

together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

b)  $V^2$ ,  $W^2$  and  $W''$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ , -

5  $\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxy carbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus;

10

c)  $Z'$  is selected from the group of  $-\text{OH}$ ,  $-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{OCO}_2\text{R}^3$ , and  $-\text{OC}(\text{O})\text{SR}^3$ ;

$D'$  is -H;

$D''$  is selected from the group of -H, alkyl,  $-\text{OR}^2$ ,  $-\text{OH}$ , and  $-\text{OC}(\text{O})\text{R}^3$ ;

15

each  $W^3$  is independently selected from the group consisting of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$p$  is an integer 2 or 3;

with the provisos that:

a)  $V$ ,  $Z$ ,  $W$ ,  $W'$  are not all -H and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all -H; and

20

b) both Y groups are not  $-\text{NR}^6$ ;

$R^2$  is selected from the group consisting of  $R^3$  and -H;

$R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

$R^6$  is selected from the group consisting of -H, and lower alkyl.

25

28. The compounds of claim 1 wherein one Y is -O-, and  $R^1$  is optionally substituted aryl; and the other Y is  $-\text{NR}^6$ -, where  $R^1$  attached to said  $-\text{NR}^6$ - is selected from the group consisting of  $-\text{C}(\text{R}^4)_2\text{C}(\text{O})\text{OR}^3$ , and  $-\text{C}(\text{R}^2)_2\text{C}(\text{O})\text{OR}^3$ .

29. The compounds of claim 1 wherein

30

$J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of -H,

-NR<sup>4</sup><sub>2</sub>, -CONR<sup>4</sup><sub>2</sub>, -CO<sub>2</sub>R<sup>3</sup>, halo, -SO<sub>2</sub>NR<sup>4</sup><sub>2</sub>, lower alkyl, lower alkenyl, lower alkylenearyl, lower alkynyl, lower perhaloalkyl, lower haloalkyl, lower aryl, lower alkylene-OH, -OR<sup>11</sup>, -CR<sup>2</sup><sub>2</sub>NR<sup>4</sup><sub>2</sub>, -CN, -C(S)NR<sup>4</sup><sub>2</sub>, -OR<sup>2</sup>, -SR<sup>2</sup>, -N<sub>3</sub>, -NO<sub>2</sub>, -NHC(S)NR<sup>4</sup><sub>2</sub>, -NR<sup>18</sup>COR<sup>2</sup>, -CR<sup>2</sup><sub>2</sub>CN;

5 L is selected from the group consisting of

i) 2,5-furanyl, 2,5-thienyl, 1,3-phenyl, 2,6-pyridyl, 2,5-oxazolyl, 5,2-oxazolyl, 2,4-oxazolyl, 4,2-oxazolyl, 2,4-imidazolyl, 2,6-pyrimidinyl, 2,6-pyrazinyl;

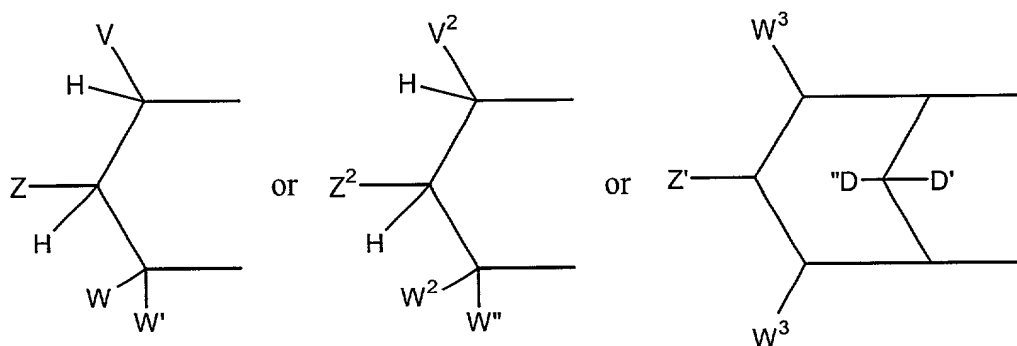
ii) 1,2-ethynyl; and

10 iii) a linking group having 3 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of alkylene-carbonylamino-, -alkylene-aminocarbonyl-, -alkylene-oxycarbonyl-, and  
15 -alkylene-oxyalkylene-;

when both Y groups are -O-, then R<sup>1</sup> is independently selected from the group consisting of optionally substituted aryl, optionally substituted benzyl, -C(R<sup>2</sup>)<sub>2</sub>OC(O)R<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>OC(O)OR<sup>3</sup>, and -H; or

when one Y is -O-, then R<sup>1</sup> attached to -O- is optionally substituted aryl; and  
20 the other Y is -NR<sup>6</sup>-, then R<sup>1</sup> attached to -NR<sup>6</sup>- is selected from the group consisting of -C(R<sup>4</sup>)<sub>2</sub>C(O)OR<sup>3</sup>, and -C(R<sup>2</sup>)<sub>2</sub>C(O)OR<sup>3</sup>; or

when Y is -O- or -NR<sup>6</sup>-, then together R<sup>1</sup> and R<sup>1</sup> are



wherein

a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

5 Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p-\text{OR}^{19}$ , and  $-(\text{CH}_2)_p-\text{SR}^{19}$ ;

or

10 together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

15

W and W' are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;

or

together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

20

b) V<sup>2</sup>, W<sup>2</sup> and W'' are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group  
5 containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus;

c)  $Z'$  is selected from the group of  $-\text{OH}$ ,  $-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{OCO}_2\text{R}^3$ , and  
10  $-\text{OC}(\text{O})\text{SR}^3$ ;

$D'$  is  $-\text{H}$ ;

$D''$  is selected from the group of  $-\text{H}$ , alkyl,  $-\text{OR}^2$ ,  $-\text{OH}$ , and  $-\text{OC}(\text{O})\text{R}^3$ ;

each  $W^3$  is independently selected from the group consisting of  $-\text{H}$ , alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

15  $p$  is an integer 2 or 3;

with the provisos that:

a)  $V$ ,  $Z$ ,  $W$ ,  $W'$  are not all  $-\text{H}$  and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all  $-\text{H}$ ; and alicyclic; and

b) both Y groups are not  $-\text{NR}^6-$ ;

20  $R^2$  is selected from the group consisting of  $R^3$  and  $-\text{H}$ ;

$R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

$R^6$  is selected from the group consisting of  $-\text{H}$ , and lower alkyl.

30. The compounds of claim 2 wherein  $R^5$  is substituted phenyl;

25 L is furan-2,5-diyl;  $J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of  $-\text{OR}^3$ ,  $-\text{SO}_2\text{NHR}^7$ ,  $-\text{CN}$ ,  $-\text{H}$ , halo,  $-\text{NR}^4_2$ ,  $-(\text{CH}_2)_2\text{aryl}$ ,  $-(\text{CH}_2)\text{NHaryl}$ , and  $-\text{NO}_2$ ; at least one Y group is  $-\text{O}-$ .

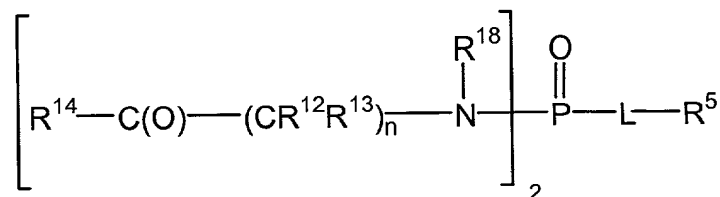
31. The compounds of claim 1 wherein



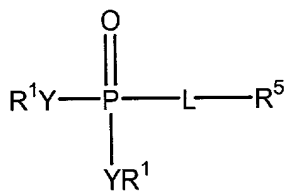
one Y is  $-\text{NR}^6-$ , and  $\text{R}^1$  attached to it is  $-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ , then the other  $\text{YR}^1$  is selected from the group consisting of  $-\text{NR}^{15}\text{R}^{16}$ ,  $-\text{OR}^7$ , and  $\text{NR}^6-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ .

32. The compounds of claim 31 wherein the other  $\text{YR}^1$  is  $-\text{OR}^7$ .

5 33. The compounds of claim 1 that are of the formula:

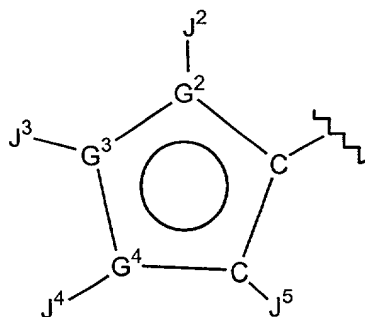


34. A method of treating a fructose-1,6-bisphosphatase dependent disease or condition in an animal which comprises administering to an animal suffering from a  
10 fructose-1,6-bisphosphatase dependent disease or condition a pharmaceutically effective amount of a compound of formula (I):



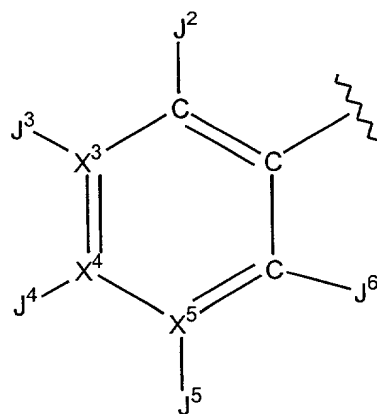
(I)

15 wherein  $\text{R}^5$  is selected from the group consisting of:



I (a)

and



I (b)

wherein:

$G^2$  is selected from the group consisting of C, O, and S;

$G^3$  and  $G^4$  are independently selected from the group consisting of C, N, O, and S;

wherein a) not more than one of  $G^2$ ,  $G^3$ , and  $G^4$  may be O, or S; b) when  $G^2$  is O or S, not more than one of  $G^3$  and  $G^4$  is N; c) at least one of  $G^2$ ,  $G^3$ , and  $G^4$  is C; and d)  $G^2$ ,  $G^3$ , and  $G^4$  are not all C;

$X^3$ ,  $X^4$ , and  $X^5$  are independently selected from the group consisting of C and N,

wherein no more than two of  $X^3$ ,  $X^4$ , and  $X^5$  may be N;

$J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of -H, -NR<sup>4</sup><sub>2</sub>, -CONR<sup>4</sup><sub>2</sub>, -CO<sub>2</sub>R<sup>3</sup>, halo, -S(O)<sub>2</sub>NR<sup>4</sup><sub>2</sub>, -S(O)R<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, alkyl, alkenyl, alkynyl, alkylenearyl, perhaloalkyl, haloalkyl, aryl, heteroaryl, alkylene-OH, -C(O)R<sup>11</sup>, -OR<sup>11</sup>, -alkylene-NR<sup>4</sup><sub>2</sub>, -alkylene-CN, -CN, -C(S)NR<sup>4</sup><sub>2</sub>, -OR<sup>2</sup>, -SR<sup>2</sup>, -N<sub>3</sub>, -NO<sub>2</sub>, -NHC(S)NR<sup>4</sup><sub>2</sub>, and -NR<sup>18</sup>COR<sup>2</sup>;

L is selected from the group consisting of:

i) a linking group having 2-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -furanyl-, -thienyl-, -pyridyl-, -oxazolyl-, -imidazolyl-, -phenyl-, -pyrimidinyl-, -pyrazinyl-, and -alkynyl-, all of which may be optionally substituted; and

ii) a linking group having 3-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -alkylenecarbonylamino-, -alkyleneaminocarbonyl-, -alkyleneoxycarbonyl-, -alkyleneoxy-, -alkylenethio-, -alkylenecarbonyloxy-, -alkylene-S(O)-, -alkylene-S(O)<sub>2</sub>-, and

-alkyleneoxyalkylene-, all of which may be optionally substituted;

Y is independently selected from the group consisting of -O-, and -NR<sup>6</sup>-;

when Y is -O-, then R<sup>1</sup> attached to -O- is independently selected from the group consisting of -H, alkyl, optionally substituted aryl, optionally substituted alicyclic where the cyclic moiety contains a carbonate or thiocarbonate, optionally substituted

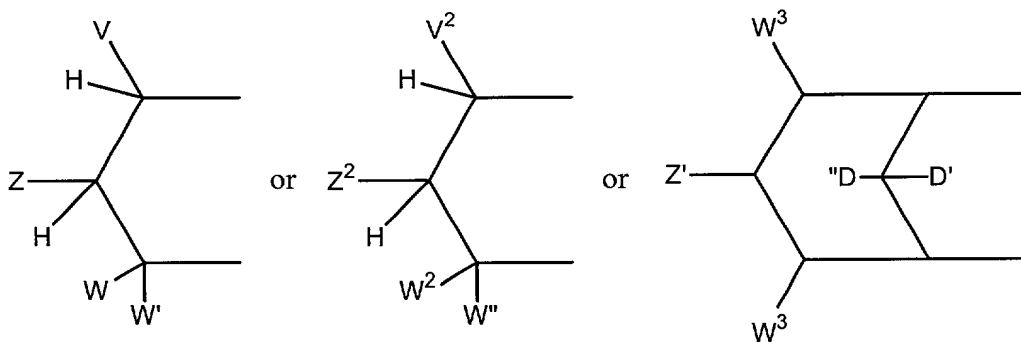
arylalkylene-,

$-\text{C}(\text{R}^2)_2\text{OC}(\text{O})\text{NR}^2_2$ ,  $-\text{NR}^2-\text{C}(\text{O})-\text{R}^3$ ,  $-\text{C}(\text{R}^2)_2-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{C}(\text{R}^2)_2-\text{O}-\text{C}(\text{O})\text{OR}^3$ ,  
 $-\text{C}(\text{R}^2)_2\text{OC}(\text{O})\text{SR}^3$ , -alkylene-S-C(O)R<sup>3</sup>, -alkylene-S-S-alkylenehydroxy, and -alkylene-S-S-alkylenehydroxy,

- when one Y is  $-\text{NR}^6-$ , and R<sup>1</sup> attached to it is  $-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ , then the other  
 5 YR<sup>1</sup> is selected from the group consisting of  $-\text{NR}^{15}\text{R}^{16}$ ,  $-\text{OR}^7$ , and  $\text{NR}^6-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ ;

or when either Y is independently selected from -O- and  $-\text{NR}^6-$ , then together R<sup>1</sup>  
 and R<sup>1</sup> are -alkylene-S-S-alkylene- to form a cyclic group, or together R<sup>1</sup> and R<sup>1</sup> are

10



wherein

- a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

- 15 Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p-\text{OR}^{19}$ , and  $-(\text{CH}_2)_p-\text{SR}^{19}$ ;  
 or

- 20 together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or  
 25 substituted heteroaryl; or

W and W' are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-R^9$ ;

or

5 together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

b)  $V^2$ ,  $W^2$  and  $W''$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

10  $Z^2$  is selected from the group of  $-CHR^2OH$ ,  $-CHR^2OC(O)R^3$ ,  $-CHR^2OC(S)R^3$ ,  $-CHR^2OCO_2R^3$ ,  $-CHR^2OC(O)SR^3$ ,  $-CHR^2OC(S)OR^3$ ,  $-CH(aryl)OH$ ,  $-CH(CH=CR^2_2)OH$ ,  $-CH(C\equiv CR^2)OH$ ,  $-SR^2$ ,  $-CH_2NHaryl$ ,  $-CH_2aryl$ ; or

15 together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus;

c)  $Z'$  is selected from the group of  $-OH$ ,  $-OC(O)R^3$ ,  $-OCO_2R^3$ , and  $-OC(O)SR^3$ ;

20  $D'$  is -H;

$D''$  is selected from the group of -H, alkyl,  $-OR^2$ ,  $-OH$ , and  $-OC(O)R^3$ ;

each  $W^3$  is independently selected from the group consisting of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

p is an integer 2 or 3;

25 with the provisos that:

a) V, Z, W, W' are not all -H and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all -H; and

$R^2$  is selected from the group consisting of  $R^3$  and -H;

$R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

each  $R^4$  is independently selected from the group consisting of -H, alkylene, -alkylenearyl and aryl, or together  $R^4$  and  $R^4$  are connected via 2-6 atoms, optionally including one heteroatom selected from the group consisting of O, N, and S;

$R^6$  is selected from the group consisting of -H, lower alkyl, acyloxyalkyl, aryl, aralkyl, alkyloxycarbonyloxyalkyl, and lower acyl, or together with  $R^{12}$  is connected via 1-4 carbon atoms to form a cyclic group;

$R^7$  is lower  $R^3$ ;

each  $R^9$  is independently selected from the group consisting of -H, alkyl, aralkyl, and alicyclic, or together  $R^9$  and  $R^9$  form a cyclic alkyl group;

$R^{11}$  is selected from the group consisting of alkyl, aryl,  $-NR^2_2$ , and  $-OR^2$ ; and

each  $R^{12}$  and  $R^{13}$  is independently selected from the group consisting of H, lower alkyl, lower aryl, lower aralkyl, all optionally substituted, or  $R^{12}$  and  $R^{13}$  together are connected via a chain of 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S, to form a cyclic group;

each  $R^{14}$  is independently selected from the group consisting of  $-OR^{17}$ ,  $-N(R^{17})_2$ ,  $-NHR^{17}$ ,  $-SR^{17}$ , and  $-NR^2OR^{20}$ ;

$R^{15}$  is selected from the group consisting of -H, lower aralkyl, lower aryl, lower aralkyl, or together with  $R^{16}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{16}$  is selected from the group consisting of  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ , -H, lower alkyl, lower aryl, lower aralkyl, or together with  $R^{15}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

each  $R^{17}$  is independently selected from the group consisting of lower alkyl, lower aryl, and lower aralkyl, or together  $R^{17}$  and  $R^{17}$  on N is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{18}$  is selected from the group consisting of -H and lower  $R^3$ ;

$R^{19}$  is selected from the group consisting of -H, and lower acyl;

$R^{20}$  is selected from the group consisting of -H, lower  $R^3$ , and  $-C(O)-(lower\ R^3)$ ;

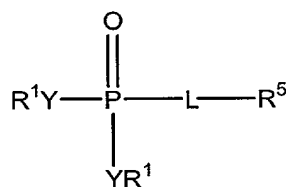
n is an integer from 1 to 3;

with the provisos that:

- 1) when  $X^3$ ,  $X^4$ , or  $X^5$  is N, then the respective  $J^3$ ,  $J^4$ , or  $J^5$  is null;
- 2) when  $G^2$ ,  $G^3$ , or  $G^4$  is O or S, then the respective  $J^2$ ,  $J^3$ , or  $J^4$  is null;
- 3) when  $G^3$  or  $G^4$  is N, then the respective  $J^3$  or  $J^4$  is not halogen or a group directly bonded to  $G^3$  or  $G^4$  via a heteroatom;
- 4) if both Y groups are  $-NR^6-$ , and  $R^1$  and  $R^1$  are not connected to form a cyclic phosphoramidate, then at least one  $R^1$  is  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ ;
- 5)  $R^1$  can be selected from the lower alkyl only when the other  $YR^1$  is  $-NR^6-C(R^{12}R^{13})_n-C(O)-R^{14}$ ;

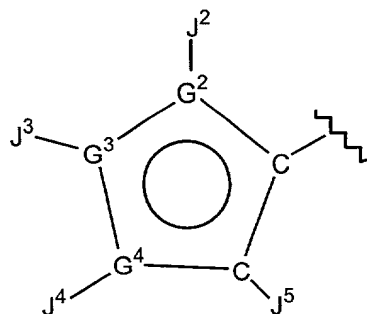
and pharmaceutically acceptable prodrugs and salts thereof.

35. A method of treating diabetes, by administering to patient in need thereof a pharmaceutically effective amount of an FBPase inhibitor of Formula I:



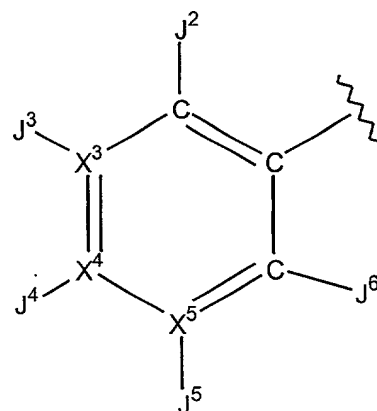
(I)

wherein  $R^5$  is selected from the group consisting of:



I (a)

and



I (b)

wherein:

$G^2$  is selected from the group consisting of C, O, and S;

$G^3$  and  $G^4$  are independently selected from the group consisting of C, N, O, and S;  
wherein a) not more than one of  $G^2$ ,  $G^3$ , and  $G^4$  may be O, or S; b) when  $G^2$  is O or S, not more than one of  $G^3$  and  $G^4$  is N; c) at least one of  $G^2$ ,  $G^3$ , and  $G^4$  is C; and d)  $G^2$ ,  $G^3$ , and  $G^4$  are not all C;

5  $X^3$ ,  $X^4$ , and  $X^5$  are independently selected from the group consisting of C and N,  
wherein no more than two of  $X^3$ ,  $X^4$ , and  $X^5$  may be N;

$J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of -H,  
-NR<sup>4</sup><sub>2</sub>, -CONR<sup>4</sup><sub>2</sub>, -CO<sub>2</sub>R<sup>3</sup>, halo, -S(O)<sub>2</sub>NR<sup>4</sup><sub>2</sub>, -S(O)R<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, alkyl, alkenyl, alkynyl,  
alkylenearyl, perhaloalkyl, haloalkyl, aryl, heteroaryl, alkylene-OH, -C(O)R<sup>11</sup>, -OR<sup>11</sup>, -  
10 alkylene-NR<sup>4</sup><sub>2</sub>, -alkylene-CN, -CN, -C(S)NR<sup>4</sup><sub>2</sub>, -OR<sup>2</sup>, -SR<sup>2</sup>, -N<sub>3</sub>, -NO<sub>2</sub>, -NHC(S)NR<sup>4</sup><sub>2</sub>, and  
-NR<sup>18</sup>COR<sup>2</sup>;

L is selected from the group consisting of:

i) a linking group having 2-4 atoms measured by the fewest number of atoms  
connecting the carbon of the aromatic ring and the phosphorus atom and is selected from  
15 the group consisting of -furanyl-, -thienyl-, -pyridyl-, -oxazolyl-, -imidazolyl-, -phenyl-,  
-pyrimidinyl-, -pyrazinyl-, and -alkynyl-, all of which may be optionally substituted; and

ii) a linking group having 3-4 atoms measured by the fewest number of atoms  
connecting the carbon of the aromatic ring and the phosphorus atom and is selected from  
the group consisting of -alkylenecarbonylamino-, -alkyleneaminocarbonyl-, -  
20 alkyleneoxycarbonyl-, -alkyleneoxy-, -alkylenethio-, -alkylenecarbonyloxy-, -alkylene-  
S(O)-, -alkylene-S(O)<sub>2</sub>-, and -alkyleneoxyalkylene-, all of which may be optionally  
substituted;

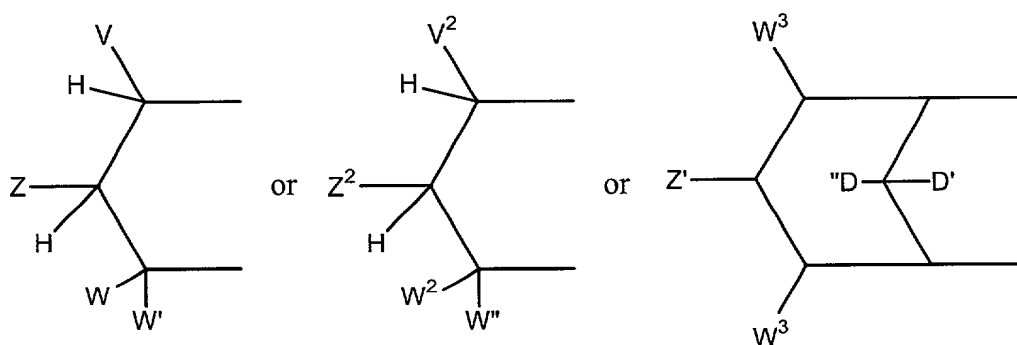
Y is independently selected from the group consisting of -O-, and -NR<sup>6</sup>-;

when Y is -O-, then R<sup>1</sup> attached to -O- is independently selected from the group  
25 consisting of -H, alkyl, optionally substituted aryl, optionally substituted alicyclic where  
the cyclic moiety contains a carbonate or thiocarbonate, optionally substituted  
arylalkylene-,

-C(R<sup>2</sup>)<sub>2</sub>OC(O)NR<sup>2</sup><sub>2</sub>, -NR<sup>2</sup>-C(O)-R<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>-OC(O)R<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>-O-C(O)OR<sup>3</sup>,  
-C(R<sup>2</sup>)<sub>2</sub>OC(O)SR<sup>3</sup>, -alkylene-S-C(O)R<sup>3</sup>, -alkylene-S-S-alkylenehydroxy, and -alkylene-S-  
30 S-S-alkylenehydroxy,

when one Y is  $-\text{NR}^6-$ , and  $\text{R}^1$  attached to it is  $-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ , then the other  $\text{YR}^1$  is selected from the group consisting of  $-\text{NR}^{15}\text{R}^{16}$ ,  $-\text{OR}^7$ , and  $\text{NR}^6-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ ;

or when either Y is independently selected from  $-\text{O}-$  and  $-\text{NR}^6-$ , then together  $\text{R}^1$  and  $\text{R}^1$  are -alkylene-S-S-alkylene- to form a cyclic group, or together  $\text{R}^1$  and  $\text{R}^1$  are



wherein

10 a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p-\text{OR}^{19}$ , and  $-(\text{CH}_2)_p-\text{SR}^{19}$ ;

15 or

together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

20 together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

W and W' are independently selected from the group of  $-\text{H}$ , alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;

25 or



together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

- 5 b)  $V^2$ ,  $W^2$  and  $W''$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

- 10 together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus;

- 15 c)  $Z'$  is selected from the group of  $-\text{OH}$ ,  $-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{OCO}_2\text{R}^3$ , and  $-\text{OC}(\text{O})\text{SR}^3$ ;

$D'$  is -H;

$D''$  is selected from the group of -H, alkyl,  $-\text{OR}^2$ ,  $-\text{OH}$ , and  $-\text{OC}(\text{O})\text{R}^3$ ;

- each  $W^3$  is independently selected from the group consisting of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;
- 20  $p$  is an integer 2 or 3;

with the provisos that:

- a) V, Z, W, W' are not all -H and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all -H; and

$R^2$  is selected from the group consisting of  $R^3$  and -H;

- 25  $R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

each  $R^4$  is independently selected from the group consisting of -H, alkylene, -alkylenearyl and aryl, or together  $R^4$  and  $R^4$  are connected via 2-6 atoms, optionally including one heteroatom selected from the group consisting of O, N, and S;

$R^6$  is selected from the group consisting of -H, lower alkyl, acyloxyalkyl, aryl, aralkyl, alkyloxycarbonyloxyalkyl, and lower acyl, or together with  $R^{12}$  is connected via 1-4 carbon atoms to form a cyclic group;

$R^7$  is lower  $R^3$ ;

5 each  $R^9$  is independently selected from the group consisting of -H, alkyl, aralkyl, and alicyclic, or together  $R^9$  and  $R^9$  form a cyclic alkyl group;

$R^{11}$  is selected from the group consisting of alkyl, aryl,  $-NR^2_2$ , and  $-OR^2$ ; and

10 each  $R^{12}$  and  $R^{13}$  is independently selected from the group consisting of H, lower alkyl, lower aryl, lower aralkyl, all optionally substituted, or  $R^{12}$  and  $R^{13}$  together are connected via a chain of 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S, to form a cyclic group;

each  $R^{14}$  is independently selected from the group consisting of  $-OR^{17}$ ,  $-N(R^{17})_2$ ,  $-NHR^{17}$ ,  $-SR^{17}$ , and  $-NR^2OR^{20}$ ;

15  $R^{15}$  is selected from the group consisting of -H, lower aralkyl, lower aryl, lower aralkyl, or together with  $R^{16}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{16}$  is selected from the group consisting of  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ , -H, lower alkyl, lower aryl, lower aralkyl, or together with  $R^{15}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

20 each  $R^{17}$  is independently selected from the group consisting of lower alkyl, lower aryl, and lower aralkyl, or together  $R^{17}$  and  $R^{17}$  on N is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{18}$  is selected from the group consisting of -H and lower  $R^3$ ;

$R^{19}$  is selected from the group consisting of -H, and lower acyl;

25  $R^{20}$  is selected from the group consisting of -H, lower  $R^3$ , and  $-C(O)-(lower\ R^3)$ ;  
 $n$  is an integer from 1 to 3;

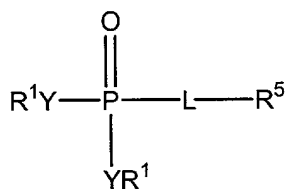
with the provisos that:

- 1) when  $X^3$ ,  $X^4$ , or  $X^5$  is N, then the respective  $J^3$ ,  $J^4$ , or  $J^5$  is null;
- 2) when  $G^2$ ,  $G^3$ , or  $G^4$  is O or S, then the respective  $J^2$ ,  $J^3$ , or  $J^4$  is null;

- 3) when  $G^3$  or  $G^4$  is N, then the respective  $J^3$  or  $J^4$  is not halogen or a group directly bonded to  $G^3$  or  $G^4$  via a heteroatom;
- 4) if both Y groups are  $-NR^6-$ , and  $R^1$  and  $R^1$  are not connected to form a cyclic phosphoramidate, then at least one  $R^1$  is  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ ;
- 5)  $R^1$  can be selected from the lower alkyl only when the other  $YR^1$  is  $-NR^6-C(R^{12}R^{13})_n-C(O)-R^{14}$ ;

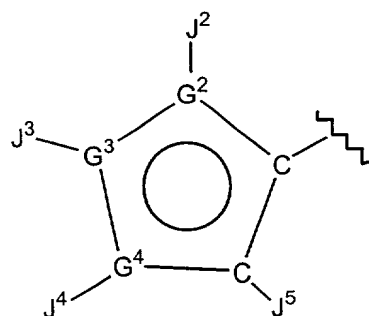
and pharmaceutically acceptable prodrugs and salts thereof.

36. A method of treating glycogen storage diseases, by administering to a patient in need thereof a pharmaceutically effective amount of an FBPase inhibitor of formula I:



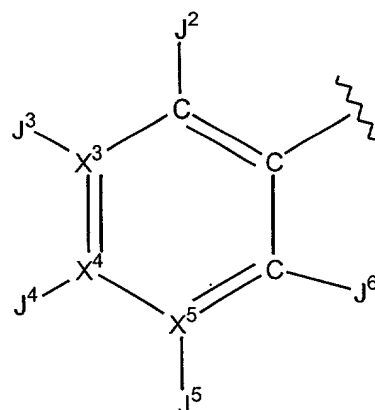
(I)

wherein  $R^5$  is selected from the group consisting of:



I (a)

and



I (b)

wherein:

$G^2$  is selected from the group consisting of C, O, and S;

$G^3$  and  $G^4$  are independently selected from the group consisting of C, N, O, and S;  
wherein a) not more than one of  $G^2$ ,  $G^3$ , and  $G^4$  may be O, or S; b) when  $G^2$  is O or S, not more than one of  $G^3$  and  $G^4$  is N; c) at least one of  $G^2$ ,  $G^3$ , and  $G^4$  is C; and d)  $G^2$ ,  $G^3$ , and  $G^4$  are not all C;

5  $X^3$ ,  $X^4$ , and  $X^5$  are independently selected from the group consisting of C and N, wherein no more than two of  $X^3$ ,  $X^4$ , and  $X^5$  may be N;

$J^2$ ,  $J^3$ ,  $J^4$ ,  $J^5$ , and  $J^6$  are independently selected from the group consisting of -H, -NR<sup>4</sup><sub>2</sub>, -CONR<sup>4</sup><sub>2</sub>, -CO<sub>2</sub>R<sup>3</sup>, halo, -S(O)<sub>2</sub>NR<sup>4</sup><sub>2</sub>, -S(O)R<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, alkyl, alkenyl, alkynyl, alkylenearyl, perhaloalkyl, haloalkyl, aryl, heteroaryl, alkylene-OH, -C(O)R<sup>11</sup>, -OR<sup>11</sup>, -alkylene-NR<sup>4</sup><sub>2</sub>, -alkylene-CN, -CN, -C(S)NR<sup>4</sup><sub>2</sub>, -OR<sup>2</sup>, -SR<sup>2</sup>, -N<sub>3</sub>, -NO<sub>2</sub>, -NHC(S)NR<sup>4</sup><sub>2</sub>, and -NR<sup>18</sup>COR<sup>2</sup>;

10

L is selected from the group consisting of:

i) a linking group having 2-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -furanyl-, -thienyl-, -pyridyl-, -oxazolyl-, -imidazolyl-, -phenyl-, -pyrimidinyl-, -pyrazinyl-, and -alkynyl-, all of which may be optionally substituted; and

15

ii) a linking group having 3-4 atoms measured by the fewest number of atoms connecting the carbon of the aromatic ring and the phosphorus atom and is selected from the group consisting of -alkylenecarbonylamino-, -alkyleneaminocarbonyl-, -alkyleneoxycarbonyl-, -alkyleneoxy-, -alkylenethio-, -alkylenecarbonyloxy-, -alkylene-S(O)-, -alkylene-S(O)<sub>2</sub>-, and -alkyleneoxyalkylene-, all of which may be optionally substituted;

20

Y is independently selected from the group consisting of -O-, and -NR<sup>6</sup>-;

when Y is -O-, then R<sup>1</sup> attached to -O- is independently selected from the group consisting of -H, alkyl, optionally substituted aryl, optionally substituted alicyclic where the cyclic moiety contains a carbonate or thiocarbonate, optionally substituted arylalkylene-,

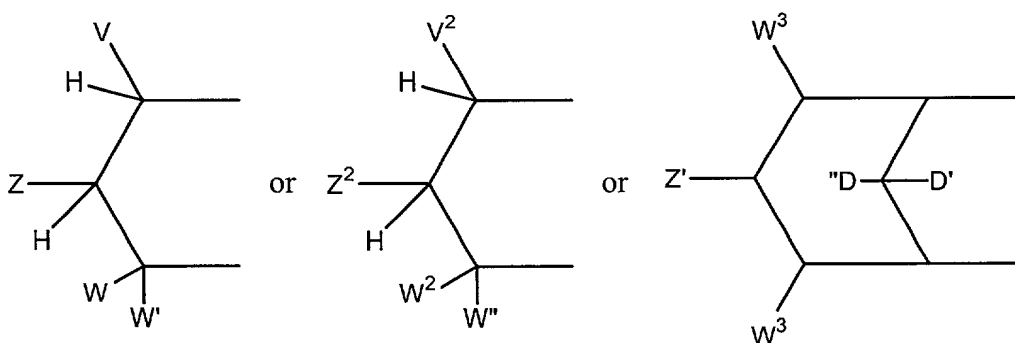
25

-C(R<sup>2</sup>)<sub>2</sub>OC(O)NR<sup>2</sup><sub>2</sub>, -NR<sup>2</sup>-C(O)-R<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>-OC(O)R<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>-O-C(O)OR<sup>3</sup>, -C(R<sup>2</sup>)<sub>2</sub>OC(O)SR<sup>3</sup>, -alkylene-S-C(O)R<sup>3</sup>, -alkylene-S-S-alkylenehydroxy, and -alkylene-S-S-S-alkylenehydroxy,

30

when one Y is  $-\text{NR}^6-$ , and  $\text{R}^1$  attached to it is  $-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ , then the other  $\text{YR}^1$  is selected from the group consisting of  $-\text{NR}^{15}\text{R}^{16}$ ,  $-\text{OR}^7$ , and  $\text{NR}^6-(\text{CR}^{12}\text{R}^{13})_n-\text{C}(\text{O})-\text{R}^{14}$ ;

or when either Y is independently selected from  $-\text{O}-$  and  $-\text{NR}^6-$ , then together  $\text{R}^1$  and  $\text{R}^1$  are -alkylene-S-S-alkylene- to form a cyclic group, or together  $\text{R}^1$  and  $\text{R}^1$  are



wherein

a) V is selected from the group of aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkynyl and 1-alkenyl;

Z is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{OR}^2$ ,  $-\text{SR}^2$ ,  $-\text{CHR}^2\text{N}_3$ ,  $-\text{CH}_2\text{aryl}$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{R}^2$ ,  $-\text{NR}^2_2$ ,  $-\text{OCOR}^3$ ,  $-\text{OCO}_2\text{R}^3$ ,  $-\text{SCOR}^3$ ,  $-\text{SCO}_2\text{R}^3$ ,  $-\text{NHCOR}^2$ ,  $-\text{NHCO}_2\text{R}^3$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-(\text{CH}_2)_p-\text{OR}^{19}$ , and  $-(\text{CH}_2)_p-\text{SR}^{19}$ ;

or

together V and Z are connected via an additional 3-5 atoms to form a cyclic group, optionally containing 1 heteroatom, said cyclic group is fused to an aryl group at the beta and gamma position to the Y adjacent to V; or

together Z and W are connected via an additional 3-5 atoms to form a cyclic group, optionally containing one heteroatom, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl; or

W and W' are independently selected from the group of  $-\text{H}$ , alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl and 1-alkynyl and  $-\text{R}^9$ ;

or

together W and W' are connected via an additional 2-5 atoms to form a cyclic group, optionally containing 0-2 heteroatoms, and V must be aryl, substituted aryl, heteroaryl, or substituted heteroaryl;

- 5 b)  $V^2$ ,  $W^2$  and  $W''$  are independently selected from the group of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

$Z^2$  is selected from the group of  $-\text{CHR}^2\text{OH}$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{R}^3$ ,  $-\text{CHR}^2\text{OCO}_2\text{R}^3$ ,  $-\text{CHR}^2\text{OC}(\text{O})\text{SR}^3$ ,  $-\text{CHR}^2\text{OC}(\text{S})\text{OR}^3$ ,  $-\text{CH}(\text{aryl})\text{OH}$ ,  $-\text{CH}(\text{CH}=\text{CR}^2_2)\text{OH}$ ,  $-\text{CH}(\text{C}\equiv\text{CR}^2)\text{OH}$ ,  $-\text{SR}^2$ ,  $-\text{CH}_2\text{NHaryl}$ ,  $-\text{CH}_2\text{aryl}$ ; or

- 10 together  $V^2$  and  $Z^2$  are connected via an additional 3-5 atoms to form a cyclic group containing 5-7 ring atoms, optionally containing 1 heteroatom, and substituted with hydroxy, acyloxy, alkyleneoxycarbonyloxy, or aryloxycarbonyloxy attached to a carbon atom that is three atoms from a Y attached to phosphorus;

- 15 c)  $Z'$  is selected from the group of  $-\text{OH}$ ,  $-\text{OC}(\text{O})\text{R}^3$ ,  $-\text{OCO}_2\text{R}^3$ , and  $-\text{OC}(\text{O})\text{SR}^3$ ;

$D'$  is -H;

$D''$  is selected from the group of -H, alkyl,  $-\text{OR}^2$ ,  $-\text{OH}$ , and  $-\text{OC}(\text{O})\text{R}^3$ ;

- 20 each  $W^3$  is independently selected from the group consisting of -H, alkyl, aralkyl, alicyclic, aryl, substituted aryl, heteroaryl, substituted heteroaryl, 1-alkenyl, and 1-alkynyl;

p is an integer 2 or 3;

with the provisos that:

- a) V, Z, W, W' are not all -H and  $V^2$ ,  $Z^2$ ,  $W^2$ ,  $W''$  are not all -H; and

$R^2$  is selected from the group consisting of  $R^3$  and -H;

- 25  $R^3$  is selected from the group consisting of alkyl, aryl, alicyclic, and aralkyl;

each  $R^4$  is independently selected from the group consisting of -H, alkylene, -alkylenearyl and aryl, or together  $R^4$  and  $R^4$  are connected via 2-6 atoms, optionally including one heteroatom selected from the group consisting of O, N, and S;

$R^6$  is selected from the group consisting of -H, lower alkyl, acyloxyalkyl, aryl, aralkyl, alkyloxycarbonyloxyalkyl, and lower acyl, or together with  $R^{12}$  is connected via 1-4 carbon atoms to form a cyclic group;

$R^7$  is lower  $R^3$ ;

5 each  $R^9$  is independently selected from the group consisting of -H, alkyl, aralkyl, and alicyclic, or together  $R^9$  and  $R^9$  form a cyclic alkyl group;

$R^{11}$  is selected from the group consisting of alkyl, aryl,  $-NR^2_2$ , and  $-OR^2$ ; and

each  $R^{12}$  and  $R^{13}$  is independently selected from the group consisting of H, lower alkyl, lower aryl, lower aralkyl, all optionally substituted, or  $R^{12}$  and  $R^{13}$  together are  
10 connected via a chain of 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S, to form a cyclic group;

each  $R^{14}$  is independently selected from the group consisting of  $-OR^{17}$ ,  $-N(R^{17})_2$ ,  $-NHR^{17}$ ,  $-SR^{17}$ , and  $-NR^2OR^{20}$ ;

$R^{15}$  is selected from the group consisting of -H, lower aralkyl, lower aryl, lower  
15 aralkyl, or together with  $R^{16}$  is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{16}$  is selected from the group consisting of  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ , -H, lower alkyl, lower aryl, lower aralkyl, or together with  $R^{15}$  is connected via 2-6 atoms, optionally  
including 1 heteroatom selected from the group consisting of O, N, and S;

20 each  $R^{17}$  is independently selected from the group consisting of lower alkyl, lower aryl, and lower aralkyl, or together  $R^{17}$  and  $R^{17}$  on N is connected via 2-6 atoms, optionally including 1 heteroatom selected from the group consisting of O, N, and S;

$R^{18}$  is selected from the group consisting of -H and lower  $R^3$ ;

$R^{19}$  is selected from the group consisting of -H, and lower acyl;

25  $R^{20}$  is selected from the group consisting of -H, lower  $R^3$ , and  $-C(O)-(lower\ R^3)$ ;  
n is an integer from 1 to 3;

with the provisos that:

- 1) when  $X^3$ ,  $X^4$ , or  $X^5$  is N, then the respective  $J^3$ ,  $J^4$ , or  $J^5$  is null;
- 2) when  $G^2$ ,  $G^3$ , or  $G^4$  is O or S, then the respective  $J^2$ ,  $J^3$ , or  $J^4$  is null;

- 3) when  $G^3$  or  $G^4$  is N, then the respective  $J^3$  or  $J^4$  is not halogen or a group directly bonded to  $G^3$  or  $G^4$  via a heteroatom;
- 4) if both Y groups are  $-NR^6-$ , and  $R^1$  and  $R^1$  are not connected to form a cyclic phosphoramidate, then at least one  $R^1$  is  $-(CR^{12}R^{13})_n-C(O)-R^{14}$ ;
- 5)  $R^1$  can be selected from the lower alkyl only when the other  $YR^1$  is  $-NR^6-C(R^{12}R^{13})_n-C(O)-R^{14}$ ;

and pharmaceutically acceptable prodrugs and salts thereof.